

WO 2005/093097

1/17

PCT/FR2005/000656

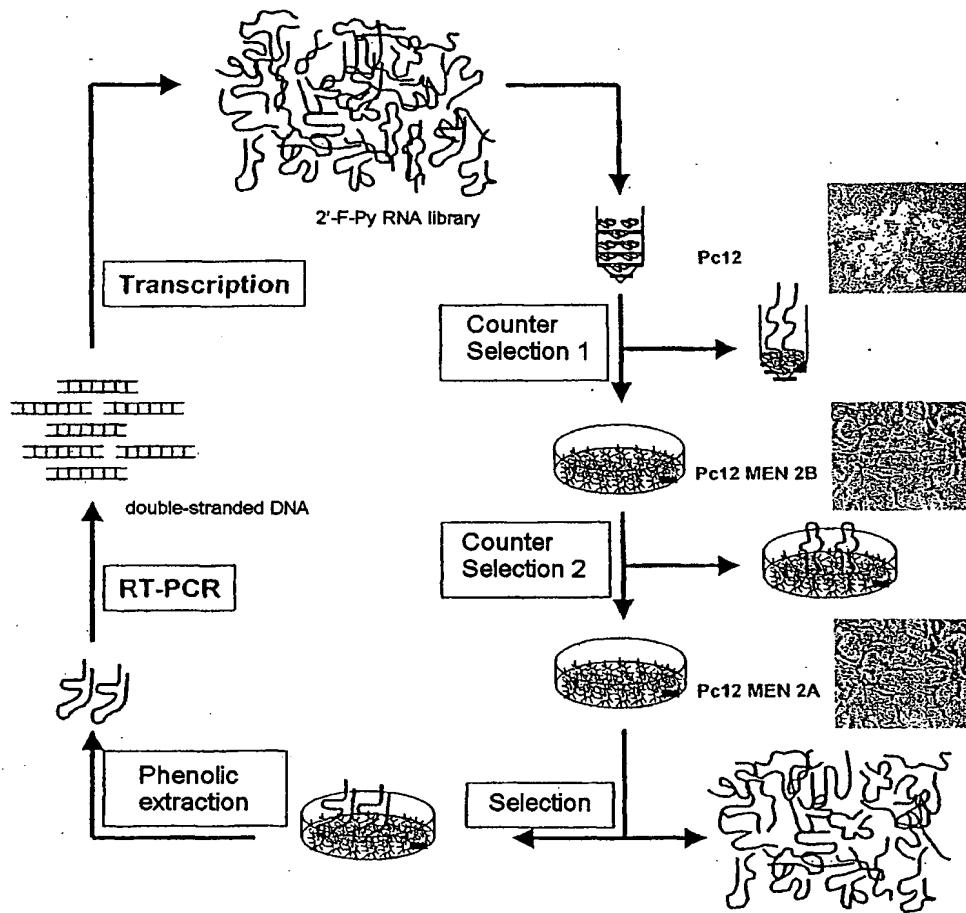


FIGURE 1

2/17

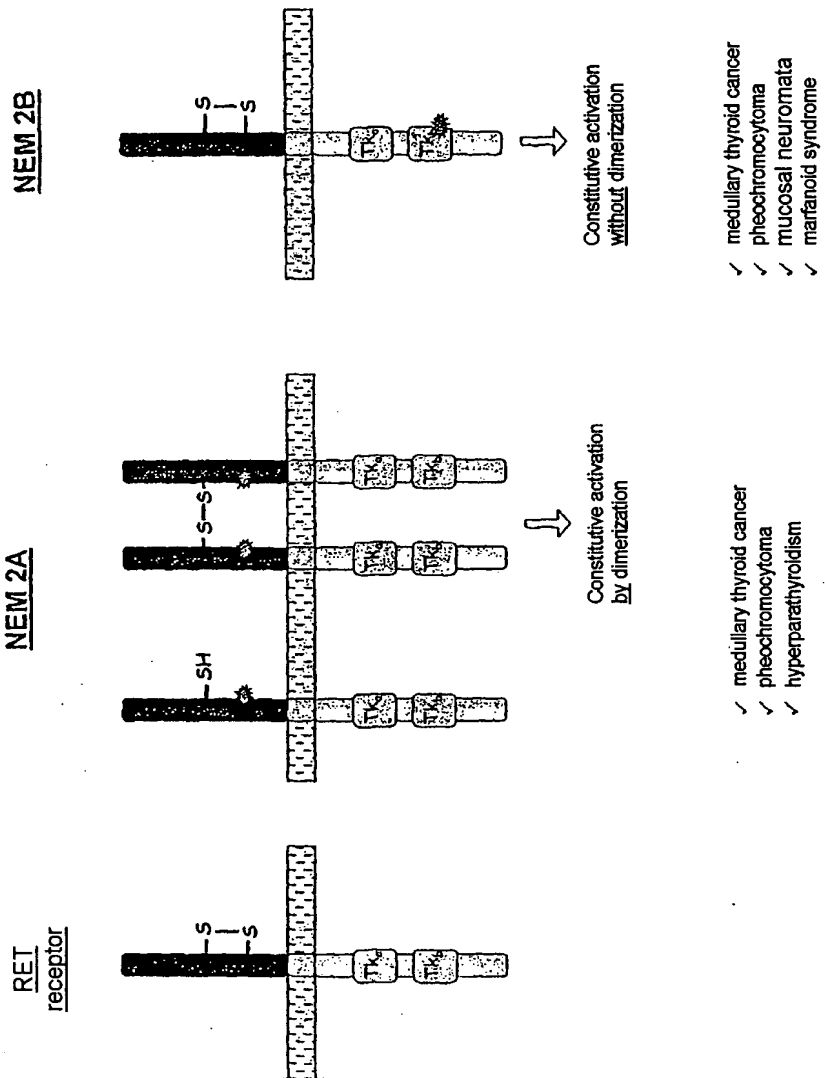
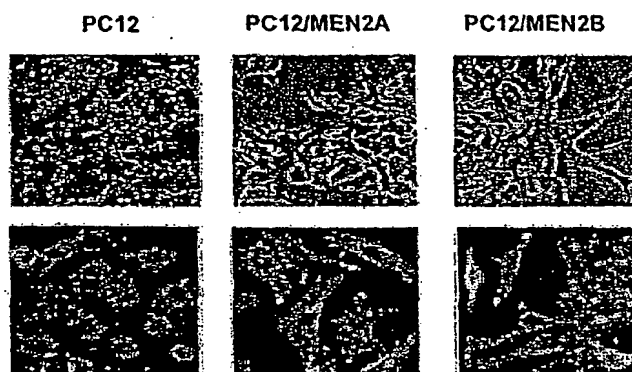


FIGURE 2

3/17



PC12 cells stably transfected with a vector for expression of the human mutated receptors Ret^{C834Y} (PC12/ MEN2A) or Ret^{M918T} (PC12/MEN2B).

FIGURE 3

4/17

Diagrammatic representation of GDNF-dependent
RET receptor activation

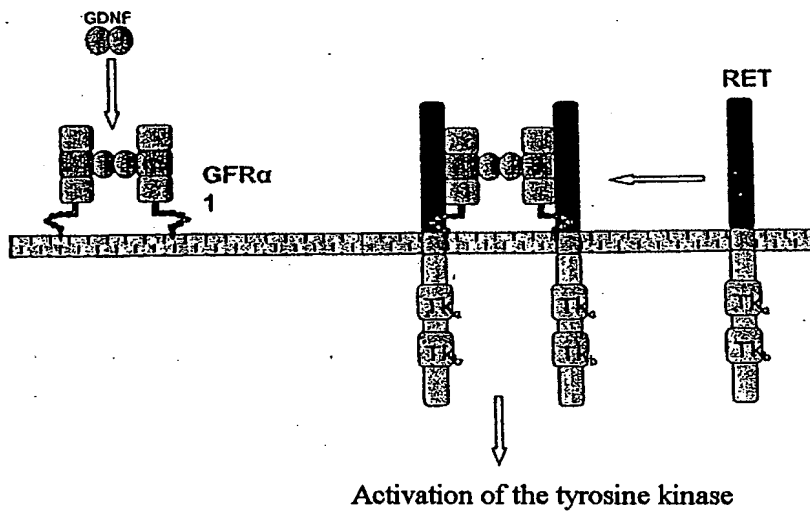


FIGURE 4

5/17

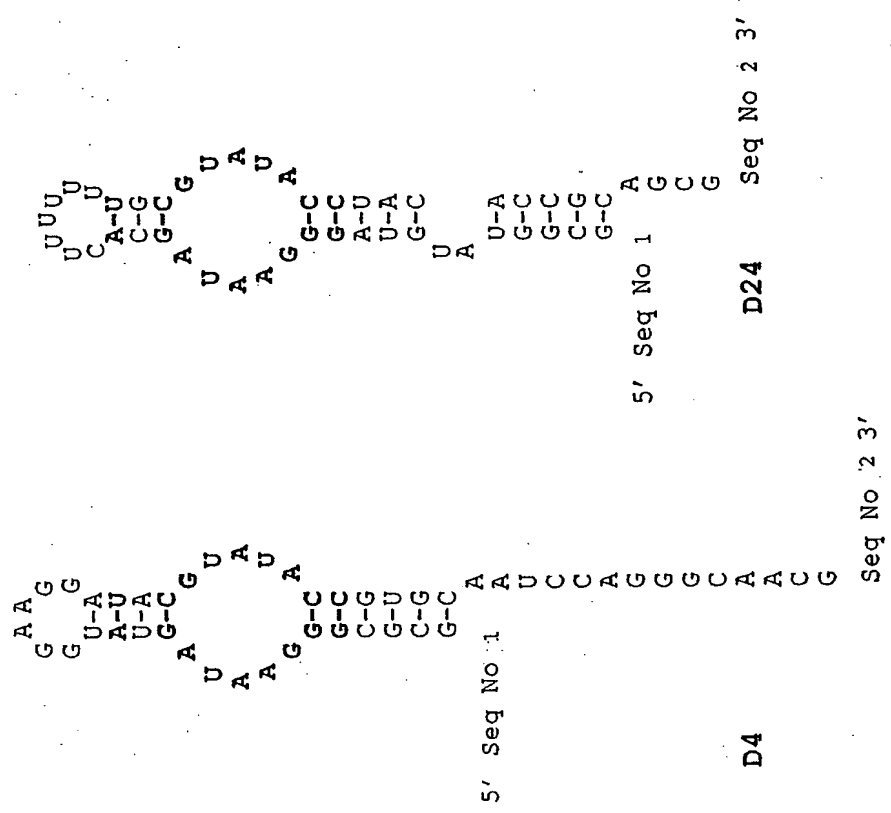


Fig. 5 A

6/17

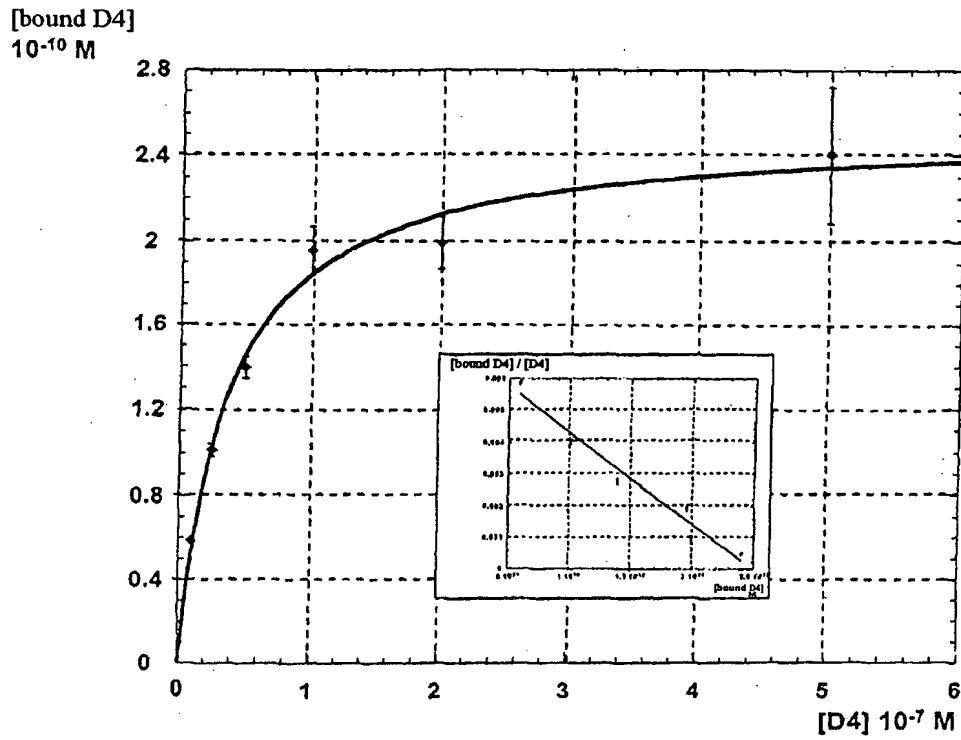
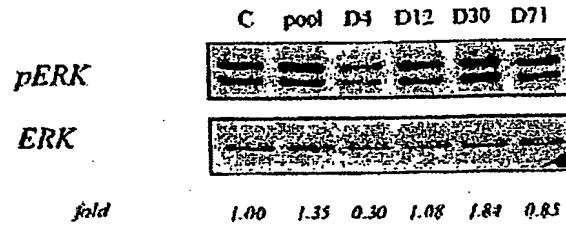


FIGURE 5B

7/17

A



B

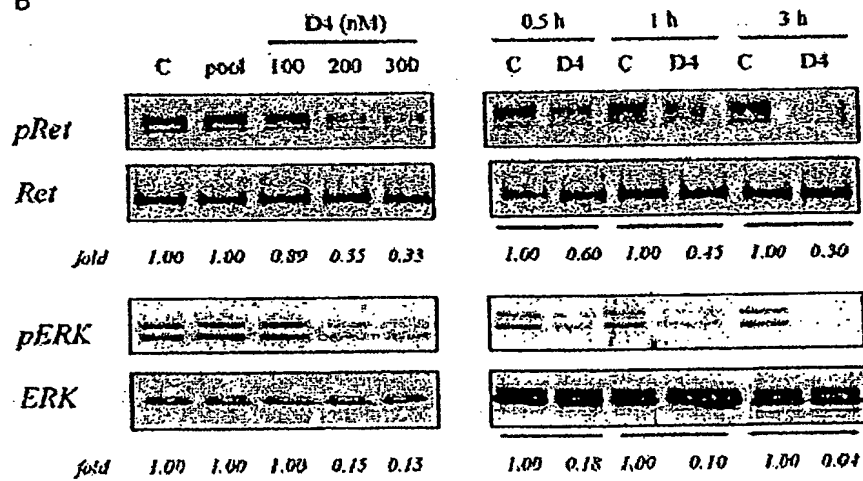


FIGURE 6

8/17

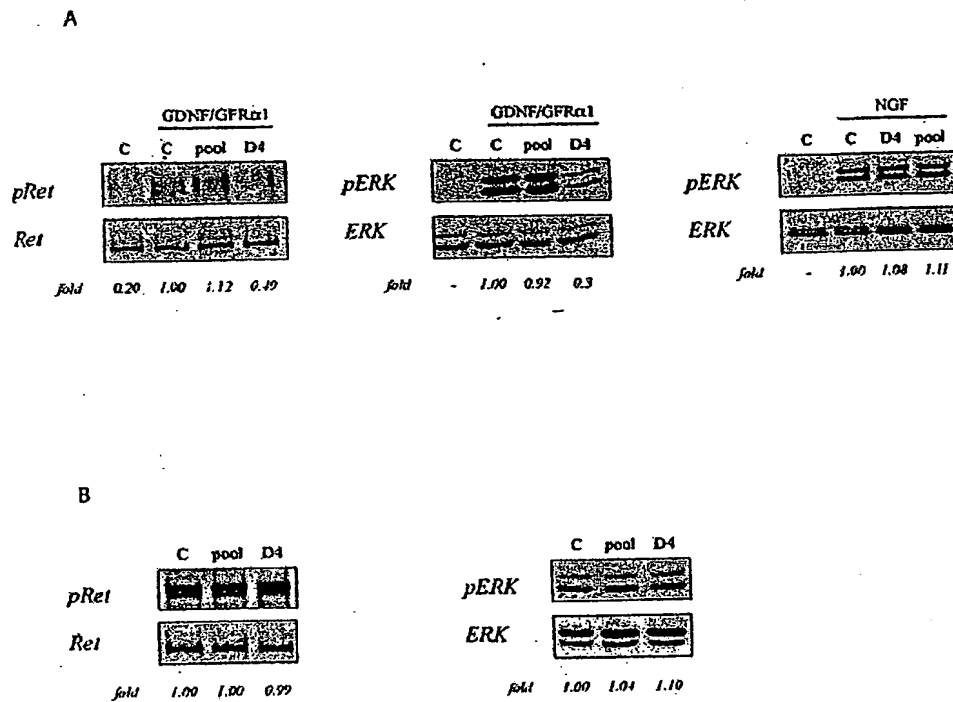


FIGURE 7

9/17

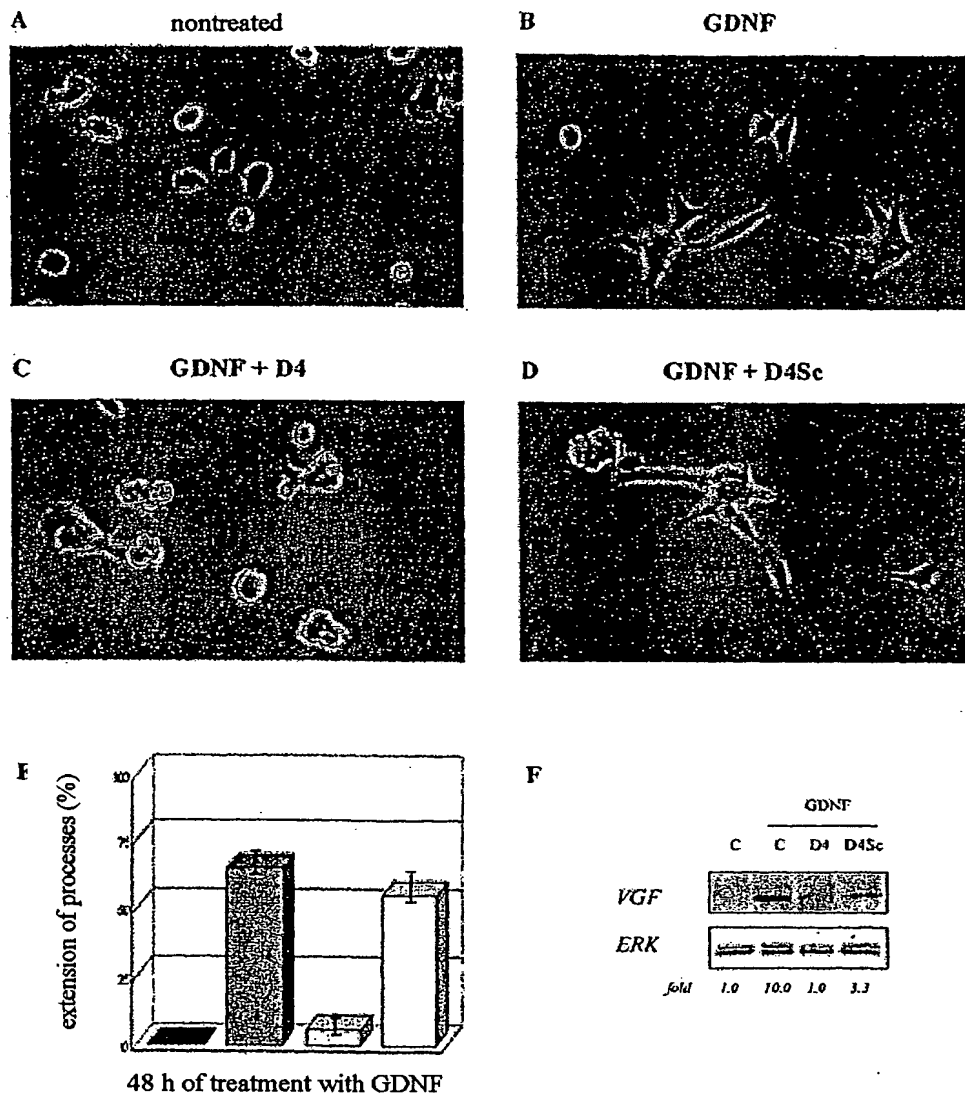
PC12- α 1/wt cells

FIGURE 8

10/17

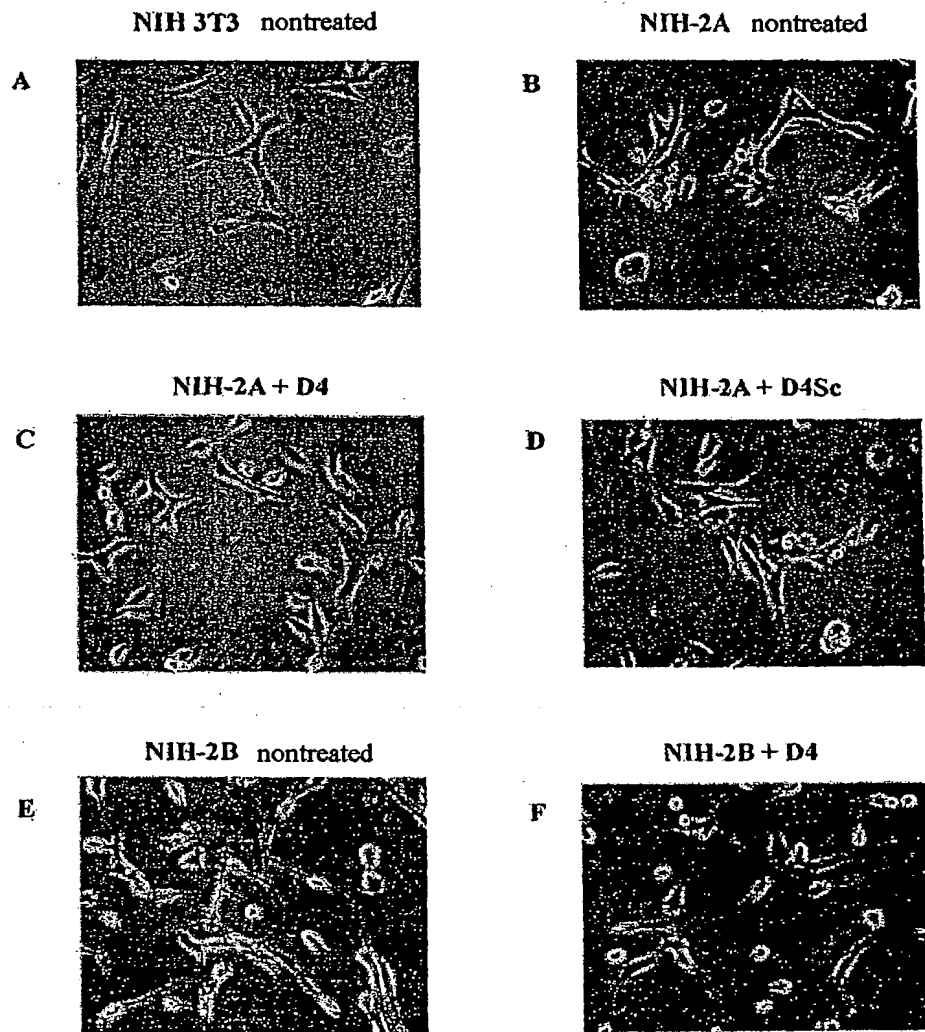
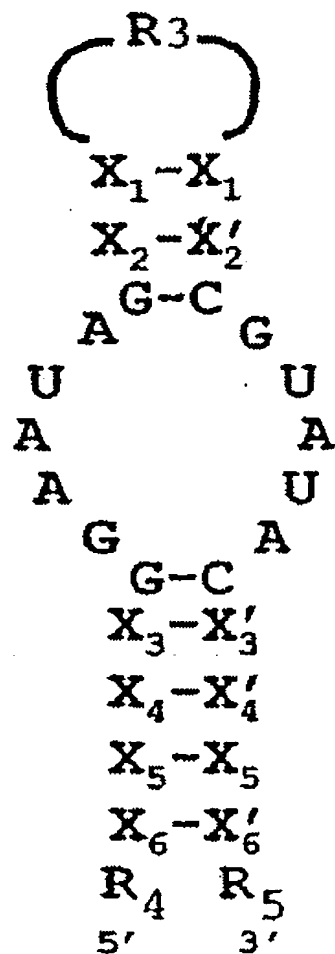


FIGURE 9

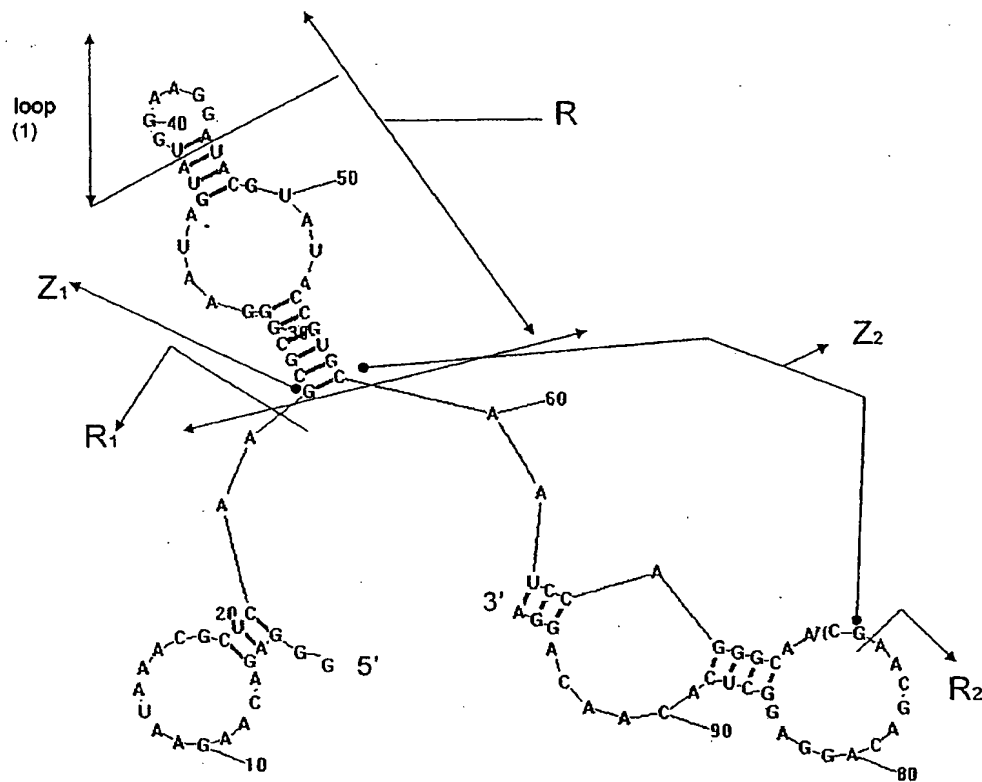
11/17

FIGURE 10



Formula II

12/17



D4 - 97nt ENERGY -17.5

FIGURE 11

5' → 3'

R₁ + Z₁ = R₄

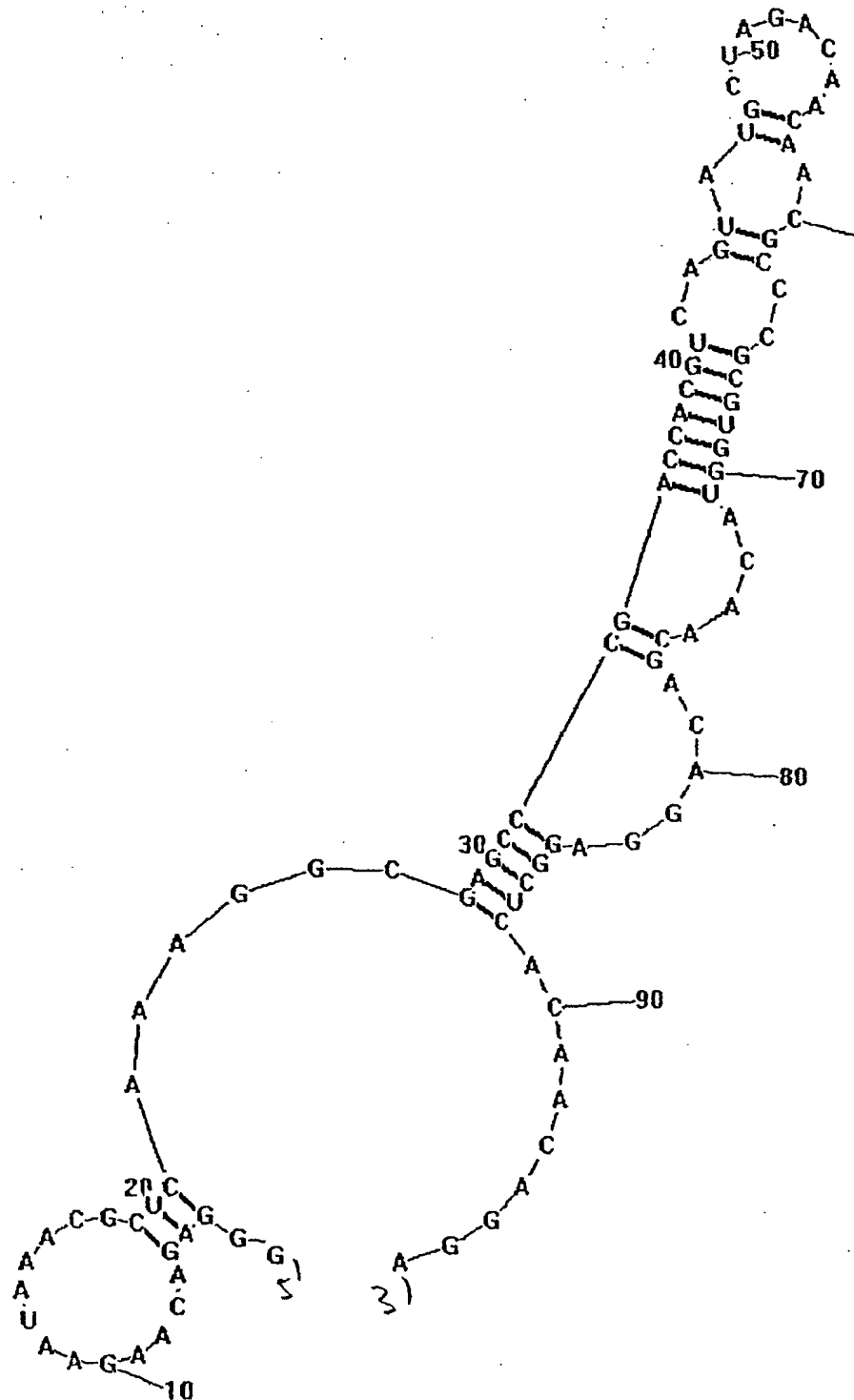
5' → 3'

R₂ + Z₂ = R₅


$$5' \rightarrow 3'$$

$$5' \rightarrow 3'$$

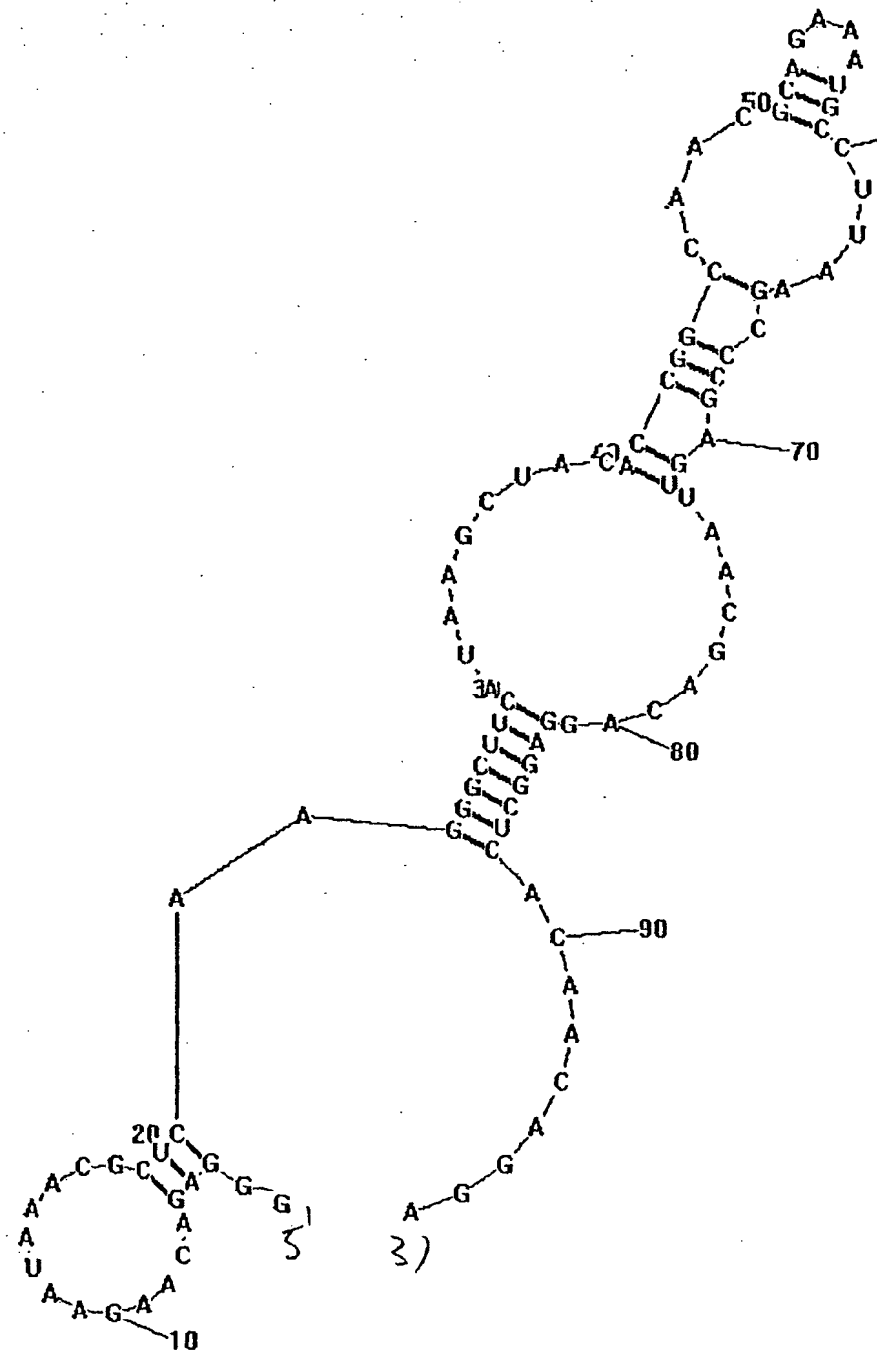
$$R_2 + Z_2 = R_5$$



D30 - 97nt ENERGY -17

FIGURE 13

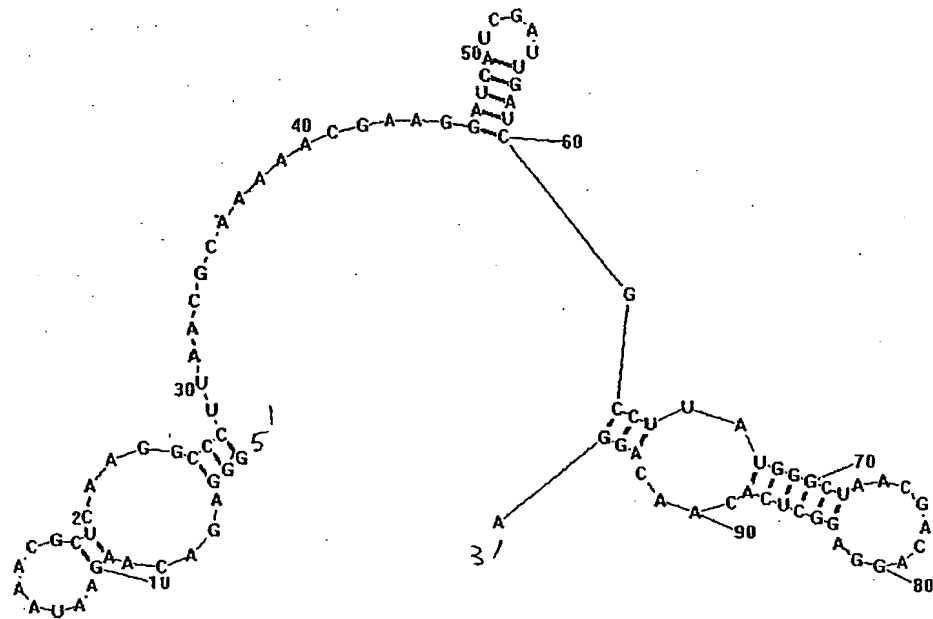
15/17



D12 - 97nt ENERGY -19.1

FIGURE 14

16/17



D71 - 96nt ENERGY -16.6

FIGURE 15

17/17

Fig. 16: Screening of aptamers which interact with RET on PC12 MEN 2A cells by competitive binding with the D4 aptamer

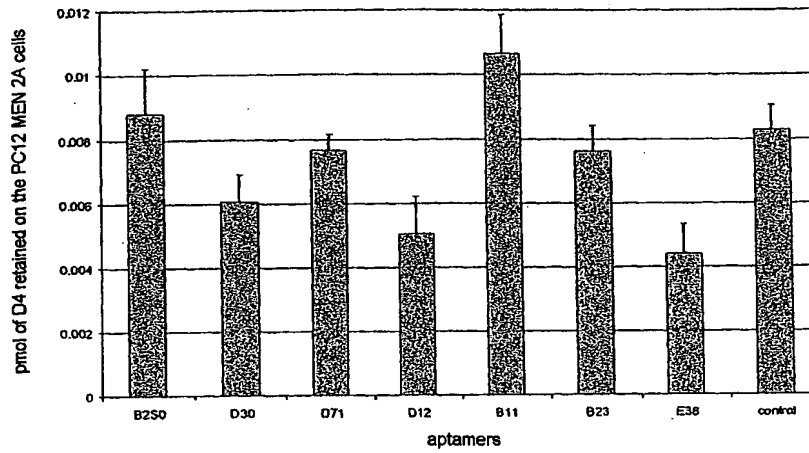


Fig. 17: Competitive binding of the E38 aptamer on PC12 MEN 2A cells in the presence of an increasing concentration of D4 aptamer

